

> S 14 AND L7
L15 81 L14 AND L7

09/516061
12/24/05

=> D 1-81 TI

=> S SEQUENCE/TI
72256 SEQUENCE/TI
37008 SEQUENCES/TI
L16 107231 SEQUENCE/TI
((SEQUENCE OR SEQUENCES)/TI)

=> S L16 AND L13
L17 477 L16 AND L13

=> S L16 AND L14
L18 136 L16 AND L14

=> D 1-136 TI

=> D 37,53 CBIB ABS

L18 ANSWER 37 OF 136 CAPLUS COPYRIGHT 2003 ACS on STN
2000:388525 Document No. 133:39121 ***Sequences*** of homologs (NL2,
NL3, and NL6) of known ligands of TIE receptor ***tyrosine***
kinase. Fong, Sherman; Ferrara, Napoleone; Goddard, Audrey;
Godowski, Paul J.; Gurney, Austin L.; Hillan, Kenneth; Williams, P. Mickey
(Genentech, Inc., USA). U.S. US 6074873 A 20000613, 50 pp.,
Cont.-in-part of U.S. Ser. No. 934,494. (English). CODEN: USXXAM.
APPLICATION: US 1998-143068 19980828. PRIORITY: US 1997-934494 19970919.
AB The invention provides protein and cDNA sequences of homologs (NL2, NL3,
and NL6) of known ligands of TIE receptor tyrosine kinase. NL3 has a
fibrinogen-like domain, has homol. with human TL-1 and human TL-2, and it
is of particular interest in this invention. The invention also relates
to the effects the provided TIE ligand homologs, esp. NL3, have on cell
proliferation, apoptosis, and angiogenesis.

L18 ANSWER 53 OF 136 CAPLUS COPYRIGHT 2003 ACS on STN
1998:187324 Document No. 128:266352 Homologous ***sequences*** in the
primary structures of ***tyrosine*** ***kinase*** receptors of the
insulin superfamily and protein-substrates 1 and 2 of the insulin
receptor. Shpakov, A. O. (I.M. Sechenov Institute of Evolutionary
Physiology and Biochemistry, Russian Academy of Sciences, Russia).
Ukrainskii Biokhimicheskii Zhurnal, 69(4), 39-48 (Russian) 1997. CODEN:
UBZHD4. ISSN: 0201-8470. Publisher: Institut Biokhimii im. A. V.
Palladina NAN Ukrainy.

AB Ligand-activated tyrosine kinase receptors of insulin superfamily peptides
can realize the signal transduction to the SH2-proteins
phosphatidylinositol 3-kinase (PI3K), protein phosphotyrosine phosphatase
(PPTP), and GRB2-adaptor protein via 2 pathways: (1) with participation of
specific proteins, the insulin receptor substrates 1 and 2 (IRS1/IRS2);
and (2) direct interaction between receptors and SH2-proteins (without
IRS-proteins). Consequently, structurally related determinants, which are
responsible for the interaction with SH2-proteins, must be present in the
receptor and IRS mols. The comparative anal. of amino acid sequences
(AAS) of human receptors for insulin, insulin-like growth factor-I and
insulin-related peptide and AAS of IRS1/IRS2 proteins allow one to
identify for the first time the long homologous regions in their primary
structures. After alignment of AAS of the regions, the sited-targets for
tyrosine phosphorylation, most important for functional activity of
tyrosine kinase receptors and IRS proteins, coincided with each other.
These results show that some homologous regions can have similar function.
Thus, the regions can be involved in coupling the receptors and
IRS-proteins with SH2-proteins, such as PI3K, PPTP, GRB2-adaptor protein.
It is also possible that the homologous regions of tyrosine kinase
receptors and IRS1/IRS2 proteins mediate the interaction between their
proteins.